SLR:dm 5585-69856-01 RCD/P101117US 474660 01/17/06

PATENT Attorney Reference No. 5585-69856-01

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of: Lu et al. Application No. 10/522,043

Filed: 10/13/2005

Confirmation No. 6728

For: POLYPEPTIDE

Examiner: Not yet assigned

Art Unit: 1614

Attorney Reference No. 5585-69856-01

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1 7 JAN 2006

Legal Staff International Division

REQUEST FOR CORRECTED OFFICIAL FILING RECEIPT

TO: PCT Legal Office

Fax No. 571-273-0459

U.S. Patent and Trademark Office (TOTAL PAGES TRANSMITTED: 8)

Two errors appear in the official Filing Receipt issued for the above-identified patent application, as follows:

- (1) The city of residence of Applicant Patricia Kuwabara should be: --Bristol--.
- (2) The title of the invention should be -- Polypeptide--.

Attached hereto is a copy of the official Filing Receipt with the requested corrections shown thereon. Also attached is a copy of the Combined Declaration and Power of Attorney signed by applicant that shows her correct city of residence and the correct title of the invention. Also attached is a copy of the first page of the specification, which shows the correct title of the invention.

Please correct the identified errors and issue a corrected official Filing Receipt.

I hereby certify that this paper and the documents referred to as being attached or enclosed herewith are being facsimile transmitted to fax number 571-273-0459 on the date shown below.

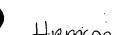
Sheree Lynn Rybak, PhD

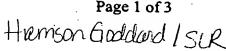
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January 17, 2006

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CONFIRMATION NO. 6728

24197 KLARQUIST SPARKMAN, LLP 121 SW SALMON STREET **SUITE 1600** PORTLAND, OR 97204

4/11/06

FILING RECEIPT

Date Mailed: 01/05/2006

Receipt is acknowledged of this regular Patent Application. It will be considered in its order and you will be notified as to the results of the examination. Be sure to provide the U.S. APPLICATION NUMBER, FILING DATE, NAME OF APPLICANT, and TITLE OF INVENTION when inquiring about this application. Fees transmitted by check or draft are subject to collection. Please verify the accuracy of the data presented on this receipt. If an error is noted on this Filing Receipt, please mail to the Commissioner for Patents P.O. Box 1450 Alexandria Va 22313-1450. Please provide a copy of this Filing Receipt with the changes noted thereon, If you received a "Notice to File Missing Parts" for this application, please submit any corrections to this Filing Receipt with your reply to the Notice. When the USPTO processes the reply to the Notice, the USPTO will generate another Filing Receipt Incorporating the requested corrections (if appropriate).

Applicant(s)

Bristol

 Xin Lu, London UNITED KINGDOM; Patricia Kuwabara, Cambridge, UNITED KINGDOM;

David Selwood, London, UNITED KINGDOM;

Power of Attorney: The patent practitioners associated with Customer Number 24197.

Domestic Priority data as claimed by applicant

This application is a 371 of PCT/GB03/04296 10/03/2003

Foreign Applications ...

UNITED KINGDOM 0223193.4 10/07/2002

UNITED KINGDOM 0306261.9 03/19/2003

Projected Publication Date: 04/13/2006

Non-Publication Request: No

Early Publication Request: No

** SMALL ENTITY **



P53 binding polypeptide e Polypeptide

Preliminary Class

514

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Almost every country has its own patent law, and a person desiring a patent in a particular country must make an application for patent in that country in accordance with its particular laws. Since the laws of many countries differ in various respects from the patent law of the United States, applicants are advised to seek guidance from specific foreign countries to ensure that patent rights are not lost prematurely.

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LUD5829 US

WDN/SLRxim 12/14/04 Combined Declarationand Power of Attorney

Express Mail Label No. EV514612324US Attorney Ref. No. 5585-69856-01

COMBINED DECLARATION AND POWER OF ATTORNEY FOR PATENT APPLICATION

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name.

joint inv	I believe I am the ventor (if plural name on the invention em			r (if only one name is li ject matter which is cla cification of which	sted below	v) or an original, for which a pater	first and It is				
	is attached hereto.										
	was filed on	_as United S	tates Patent Appl	ication No		•					
\boxtimes	was described and claimed in PCT International Application No. PCT/GB2003/004296, filed on 03 October 2003, and as amended under PCT Articles 19 on (if applicable).										
\boxtimes	and was amended				,						
	with amendments through (if applicable).										
includir	I hereby state that ng the claims, as an	I have review	wed and understa	nd the contents of the a	bove-ident	dified specification	n,				
acknow	re claims and subjected the duty to distance the duty to distance application.	ct matter in a isclose materi lication and the	application filed to distinct the distinct to that distinct as its information as the national or PC	which is material to pat inder the conditions spe- closed in the prior cope defined in 37 C.F.R. § T international filing di	ecified in 3 ending app 1.56 which are of the c	5 U.S.C. § 120 v lication, I further h occurred between continuation-in-pa	vhich cen the art				
invento States o	States of America I	isted below a	nd have also ider	SU.S.C. § 119(a)-(d) of ication(s) designating a stifled below any forcig n(s) designating at lease r baving a filing date be	n applicat	country other the ion(s) for patent	in the or				
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	0223193	.4 (Freat Britain	07 October 200	02	Claim Priority?					
٠	030626]	.9 (Great Britain	19 March 200	3	Yes No Yes No	-				
below:	I hereby claim the	benefit unde	er 35 U.S.C. § 11	%e) of any United State	es provisio	mal application(s) listed				
Application Number				Filing Date							
							-				

WDN/5LR:dm 07/22/05 356929 doc

Attorney Ref No 5585-69856-01

I hereby claim the benefit under 35 U.S.C. § 120 of any United States application(s) or § 365(c) of any PCI international application(s) designating the United States, listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States or PCI International application in the matter provided by the first paragraph of Iftle 35, United States Code, § 112, I acknowledge the duty to disclose material information as defined in Title 37, Code of Federal Regulations, § 1 56(a) which occurred between the filing date of the prior application and the national or PCI international filing date of this application:

Application Number

Filing Date

Status: patented, pending, abandoned

I hereby appoint the practitioners associated with the customer number provided below to prosecute this application, to file a corresponding international application, and to transact all business in the Patent and I rademark

Customer Number 24197

I hereby grant the law firm of Klarquist Sparkman, LLP, the power to insert on this Combined Declaration and Power of Attorney any further identification which may be necessary or desirable in order to comply with the rules of the United States Patent and Trademark Office for submitting this document.

Address all telephone calls to Sheree Lynn Rybak, Ph.D at telephone number (503) 226-7391.

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Klarquist Sparkman, LLP 121 S.W. Salmon Street, Suite 1600 Portland, OR 97204

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of I itle 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon

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Date 7th Sept

COMBINED DECLARATION AND POWER OF ATTORNEY - PAGE 2 of 3

5829 US.

WDN/SLR:dm 12/14/04 Combined Declaration and Power of Attorney

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COMBINED DECLARATION AND POWER OF ATTORNEY - PAGE 3 of 3

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WO 2004/031229

PCT/GB2003/004296

POLYPEPTIDE

The invention relates to a polypeptide, or part thereof, which inhibits the apoptotic activity of the tumour suppressor protein p53; screening methods to identify agents which interfere with the activity of said polypeptide and agents with said activity.

. Apoptosis, or programmed cell death, is a process by which multi-cellular organisms regulate cell number and differentiation. The process is regulated by factors which either induce or prevent apoptosis. Inducers of apoptosis include Bcl-2 10 family members, caspase family members and their associated factors Apaf-1 and Fadd. Caspases are synthesised as proenzymes which become activated after proteolytic cleavage. The active caspase then induces many of the morphological and biochemical changes associated with apoptosis. Mitochondria play a pivotal role in the activation process through the release of pro-apoptotic factors such as cytochrome c, AIF and Diablo. The release from mitochondria is controlled by the Bcl-2 family of proteins; (e.g. Bcl-2 and Bcl-xl inhibit release; Bax and Bak induce release).

WO9953051 discloses a cytokine dependent protein p21 which has pro-apoptotic activity. p21 is expressed in a cytokine dependent manner in myeloid/enthyroid cells. These cells are dependent on IL-3 for growth and in the absence of IL-3 the translation of p21 is induced resulting in apoptosis and cell death. p21 is a cytoplasmic protein which translocates to the outer mitochondrial membrane to induce pro-apoptotic activities.

Turnour suppressor proteins also have pro-apoptotic activities. 25

Tumour suppressor genes encode proteins which function to inhibit cell growth or division and are therefore important with respect to maintaining proliferation, growth and differentiation of normal cells. Mutations in tumour suppressor genes result in abnormal cell-cycle progression whereby the normal cell-cycle check points which arrest the cell-cycle, when, for example, DNA is damaged, are ignored and damaged